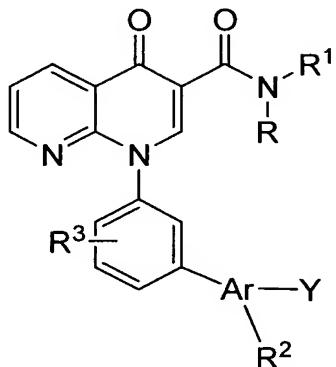


AMENDMENTS TO THE CLAIMS

This listing of claims will replace all prior versions, and listing of claims in the application.

1. (Canceled)

2. (Presently amended) A compound represented by Formula (I):



(I)

or a pharmaceutically acceptable salt thereof, wherein

Ar is phenyl, pyridyl, pyrimidyl, indolyl, quinolinyl, thienyl, pyridonyl, oxazolyl, oxadiazolyl, thiadiazolyl, or imidazolyl; or oxides thereof when Ar is a heteroaryl; Y is -COOH, or -C1-6alkyl(C1-4alkyl)_n-COOH, -C3-4cycloalkyl(C1-4alkyl)_m-COOH, wherein the -C1-6alkyl and the C3-4cycloalkyl is optionally substituted with halogen, alkoxy, hydroxy or nitrile, and the (C1-4alkyl) substituents are optionally linked to form a C3-4cycloalkyl; wherein n is 0, 1, 2, 3 or 4, m is 0, 1 or 2;

R is H or -C1-6alkyl;

R¹ is H, or -C1-6alkyl, -C3-6cycloalkyl, -C1-6alkoxy, -C2-6alkenyl, -C3-6alkynyl, heteroaryl, or heterocycle group, optionally substituted with 1-3 independent haloC1-6alkyl, -C1-6alkyl, -C1-6alkoxy, OH, amino, -(C0-6alkyl)-SO_p-(C1-6alkyl), nitro, CN, =N-O-C1-6alkyl, -O-N=C1-6alkyl, or halogen substituents, wherein p is 0, 1 or 2;

R² is H, halogen, -CN, -NO₂, -C1-6alkyl, -C3-6cycloalkyl, -O- C3-6cycloalkyl, O-C1-6alkyl, O-C3-6cycloalkyl-C1-6alkyl(C3-6cycloalkyl)(C3-6cycloalkyl), -C1-6alkoxy, phenyl, heteroaryl, heterocycle, amino, -C(O)-C1-6alkyl, -C(O)-O-C1-6alkyl, -C1-6alkyl(=N-OH), -C(N=NOH)C1-6alkyl, -C0-6alkyl(oxy)C1-6alkyl-phenyl, -SO_kNH(C0-6alkyl), or -(C0-6alkyl)-SO_k-(C1-6alkyl), wherein the phenyl, heteroaryl or heterocycle is optionally substituted with halogen, -C1-6alkyl, -C1-6alkoxy, hydroxy, amino,

or $-C(O)-O-C_1-6alkyl$, and wherein the alkyl or cycloalkyl is optionally substituted with 1-6 independently selected halogens or $-OH$, and wherein k is 0, 1, or 2;

R^3 is selected from H, halogen, CN, $-C_1-6alkyl$, $-C_3-6cycloalkyl$, nitro, $-C(O)-C_1-6alkyl$, $-C(O)-O-C_0-6alkyl$, $-SO_n'NH(C_0-6alkyl)$, or $-(C_0-6alkyl)-SO_n'-(C_1-6alkyl)$, $O-C_1-6alkyl$, $O-C_3-6cycloalkyl$, wherein n' is 0, 1, or 2 and wherein the alkyl and cycloalkyl is optionally substituted with 1-6 independently selected halogen or OH.

3 to 10. (Canceled)

11. (Original) The compound according to claim 2, or a pharmaceutically acceptable salt, wherein

Ar is pyridyl, pyrimidyl, or oxide thereof.

12. (Original) The compound according to claim 11, or a pharmaceutically acceptable salt, wherein

R^1 is $-C_1-6alkyl$ optionally substituted with 1-3 independent $-C_1-6alkyl$, $-C_1-6alkoxy$, OH, amino, $-(C_0-6alkyl)-SO_p-(C_1-6alkyl)$, nitro, CN, $=N-O-C_1-6alkyl$, $-O-N=C_1-6alkyl$, or halogen substituents.

13. (Original) The compound according to claim 11, or a pharmaceutically acceptable salt thereof, wherein

R^1 is $-C_3-6cycloalkyl$ optionally substituted with 1-3 independent $-C_1-6alkyl$, $-C_1-6alkoxy$, OH, amino, $-(C_0-6alkyl)-SO_p-(C_1-6alkyl)$, nitro, CN, $=N-O-C_1-6alkyl$, $-O-N=C_1-6alkyl$, or halogen substituents.

14. (Original) The compound according to claim 11, or a pharmaceutically acceptable salt thereof, wherein

R is hydrogen.

15. (Original) The compound according to claim 11, or a pharmaceutically acceptable salt thereof, wherein

R^2 is hydrogen or $-C_1-3alkyl$ or halogen.

16. (Original) The compound according to claim 11, or a pharmaceutically acceptable salt thereof, wherein

R^1 is $-C_3-6cycloalkyl$ optionally substituted with methyl or halo; and

R is hydrogen.

17. (Original) The compound according to claim 11, or a pharmaceutically acceptable salt thereof, wherein

R¹ is cyclopropyl optionally substituted with methyl or halo; and

R and R² are hydrogen or halogen;

R³ is hydrogen or halogen.

18. (Original) The compound according to claim 2, or a pharmaceutically acceptable salt thereof, wherein

R and R³ are hydrogen,;

R¹ is -C₃-6cycloalkyl optionally substituted with methyl or halo, or -C₁₋₃alkyl optionally substituted with 1-3 halo; and

Ar is phenyl.

19. (Cancelled)

20. (Original) The compound according to claim 2, which is 2-(trans)-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;

2-(trans)-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-3-yl}cyclopropanecarboxylic acid;

2-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-3-yl}-2-methylpropanoic acid;

2-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-4-yl}-2-methylpropanoic acid;

3-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-4-yl}-3-methylbutanoic acid;

{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-4-yl}(hydroxy)acetic acid;

1-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;

2-(cis)-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;

5-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-4-yl}-2,2-dimethyl-1,3-dioxolane-4-carboxylic acid;
1-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-3-yl}cyclopropanecarboxylic acid;
1-cyano-3-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-4-yl}-2,2-dimethylcyclopropanecarboxylic acid;
2-(trans)-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-3-fluoro-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;
(cis)-2-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-3-yl}cyclopropanecarboxylic acid;
2-(trans)-{3'-bromo-5'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;
2-(trans)-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-3-methyl-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;
2-(trans)-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-2-methyl-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;
2-(trans)-{3-chloro-3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;
2-(cis)-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-3-fluoro-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;
3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-4-carboxylic acid;
2-(trans)-{3'-[3-(morpholin-4-ylcarbonyl)-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;
2-(trans)-{3'-[4-oxo-3-({[5-(trifluoromethyl)-1,3,4-thiadiazol-2-yl]amino}carbonyl)-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;
2-(trans)-{3'-[3-({[2-(methylthio)ethyl]amino}carbonyl)-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;
2-(trans)-{3'-[3-({[2-(methylsulfonyl)ethyl]amino}carbonyl)-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;
2-(trans)-{3'-[4-oxo-3-{{(2,2,2-trifluoroethyl)amino}carbonyl}-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;

2-(trans)-(5-{3-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4*H*)-yl]phenyl}thien-2-yl)cyclopropanecarboxylic acid;
2-(trans)-{3'-[3-[(cyclopropylmethyl)amino]carbonyl]-4-oxo-1,8-naphthyridin-1(4*H*)-yl]-1,1'-biphenyl-4-yl)cyclopropanecarboxylic acid;
2-(trans)-{3'-[3-[(1-cyanocyclopropyl)amino]carbonyl]-4-oxo-1,8-naphthyridin-1(4*H*)-yl]-1,1'-biphenyl-4-yl)cyclopropanecarboxylic acid; or
3-{3'-[3-[(isopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4*H*)-yl]-1,1'-biphenyl-4-yl}-3-methylbutanoic acid.

21. (Presently amended) A compound of claim 1 which is
(+)-(trans)-2-{3-fluoro-3'-[4-oxo-3-[(2,2,2-trifluoroethyl)amino]carbonyl]-1,8-naphthyridin-1(4*H*)-yl]biphenyl-4-yl)cyclopropanecarboxylic acid;
1-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4*H*)-yl]biphenyl-4-yl}methylcyclobutanecarboxylic acid;
(trans)-2-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4*H*)-yl]biphenyl-4-yl}-2-methylcyclopropanecarboxylic acid;
(trans)-2-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4*H*)-yl]biphenyl-2-yl)cyclopropanecarboxylic acid;
3-methyl-3-{3'-[4-oxo-3-[(2,2,2-trifluoroethyl)amino]carbonyl]-1,8-naphthyridin-1(4*H*)-yl]biphenyl-4-yl}butanoic acid;
(trans)-2-{3'-[4-oxo-3-[(2,2,2-trifluoroethyl)amino]carbonyl]-1,8-naphthyridin-1(4*H*)-yl]biphenyl-2-yl)cyclopropanecarboxylic acid;
(trans)-2-{3'-[4-oxo-3-[(2,2,3,3,3-pentafluoropropyl)amino]carbonyl]-1,8-naphthyridin-1(4*H*)-yl]biphenyl-4-yl)cyclopropanecarboxylic acid;
(trans)-2-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4*H*)-yl]biphenyl-4-yl}-1-fluorocyclopropanecarboxylic acid;
(+)-(trans)-2-{3-chloro-3'-(3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4*H*)-yl]biphenyl-4-yl)cyclopropanecarboxylic acid;
(-)-(trans)-2-{3'-(4-oxo-3-[(2,2,2-trifluoroethyl)amino]carbonyl)-1,8-naphthyridin-1(4*H*)-yl]biphenyl-4-yl)cyclopropanecarboxylic acid;

(+)-(trans)-ethyl 2-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}cyclopropanecarboxylate;

(+)-(trans)-isopropyl 2-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}cyclopropanecarboxylate;

tert-butyl 3-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}-2,2-dimethylpropanoate;

3-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}-2,2-dimethylpropanoic acid;

3-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-3-yl}-2,2-dimethylpropanoic acid;

1-({3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-3-yl}methyl)cyclobutanecarboxylic acid;

3-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-2-yl}-2,2-dimethylpropanoic acid;

1-({3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-2-yl}methyl)cyclobutanecarboxylic acid;

(+)-(trans)-2-{3'-[3-[(*tert*-butylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}cyclopropanecarboxylic acid;

(+)-(trans)-2-{3'-[3-[(cyclobutylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}cyclopropanecarboxylic acid;

3-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}bicyclo[1.1.1]pentane-1-carboxylic acid;

4-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}-4-hydroxypentanoic acid;

(trans)-2-{3'-[3-{[(\square)-*cis*-(2-fluorocyclopropyl)amino]carbonyl}-4-oxo-1,8-naphthyridin-1(4H)-yl]-(+)-biphenyl-4-yl}cyclopropanecarboxylic acid;

(+)-(trans)-2-{3'-[3-{[(dicyclopropylmethyl)amino]carbonyl}-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}cyclopropanecarboxylic acid;

4-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}-2,2-dimethylbutanoic acid;

(+)-(trans)-2-{3'-[3-{[(1-hydroxycyclopropyl)amino]carbonyl}-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}cyclopropanecarboxylic acid;

(+)-(trans)-2-{3'-[4-oxo-3-{[(1-phenylcyclopropyl)amino]carbonyl}-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}cyclopropanecarboxylic acid;

4-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}-3,3-dimethylbutanoic acid;

(+)-(trans)-2-{3'-[3-{[(1-cyclopropyl-1-methylethyl)amino]carbonyl}-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}cyclopropanecarboxylic acid;

1-({3'-[4-oxo-3-{[(2,2,2-trifluoroethyl)amino]carbonyl}-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}methyl)cyclobutanecarboxylic acid;

(+)-(trans)-2-{3'-[3-{[(cyclopropylmethyl)amino]carbonyl}-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}cyclopropanecarboxylic acid;

(-)-(trans)-2-{3-fluoro-3'-[3-{[(1-hydroxycyclopropyl)amino]carbonyl}-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}cyclopropanecarboxylic acid;

(trans)-2-{3'-[4-oxo-3-{[(\square)-2,2,2-trifluoro-1-methylethyl)amino]carbonyl}-1,8-naphthyridin-1(4H)-yl]-(+)-biphenyl-4-yl}cyclopropanecarboxylic acid;

(+)-(trans)-2-{3'-[3-{[(1-methylcyclopropyl)amino]carbonyl}-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}cyclopropanecarboxylic acid;

2,2-dimethyl-4-{3'-[4-oxo-3-{[(2,2,2-trifluoroethyl)amino]carbonyl}-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}butanoic acid;

2,2-dimethyl-3-{3'-[4-oxo-3-{[(2,2,2-trifluoroethyl)amino]carbonyl}-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}propanoic acid;

(-)-(trans)-2-{3-chloro-3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}cyclopropanecarboxylic acid; or

(+)-(trans)-2-{3'-[4-oxo-3-{[(2,2,2-trifluoroethyl)amino]carbonyl}-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}cyclopropanecarboxylic acid.

22. (Presently amended) A pharmaceutical composition comprising a therapeutically effective amount of the compound according to claim 1 claim 2 or a pharmaceutically acceptable salt thereof; and a pharmaceutically acceptable carrier.

23 to 24. (Canceled).

25. (Presently amended) A method of treatment or prevention of asthma, chronic bronchitis, chronic obstructive pulmonary disease (COPD), eosinophilic granuloma, psoriasis and other benign or malignant proliferative skin diseases, endotoxic shock (and associated conditions such as laminitis and colic in horses), septic shock, ulcerative colitis, Crohn's disease, reperfusion injury of the myocardium and brain, inflammatory arthritis, osteoporosis, chronic glomerulonephritis, atopic dermatitis, urticaria, adult respiratory distress syndrome, infant respiratory distress syndrome, chronic obstructive pulmonary disease in animals, diabetes insipidus, allergic rhinitis, allergic conjunctivitis, vernal conjunctivitis, arterial restenosis, atherosclerosis, neurogenic inflammation, pain, cough, rheumatoid arthritis, ankylosing spondylitis, transplant rejection and graft versus host disease, hypersecretion of gastric acid, bacterial, fungal or viral induced sepsis or septic shock, inflammation and cytokine-mediated chronic tissue degeneration, osteoarthritis, cancer, cachexia, muscle wasting, depression, memory impairment, monopolar depression, acute and chronic neurodegenerative disorders with inflammatory components, Parkinson disease, Alzheimer's disease, spinal cord trauma, head injury, multiple sclerosis, tumour growth and cancerous invasion of normal tissues comprising the step of administering a therapeutically effective amount, or a prophylactically effective amount, of the compound according to claim 1 or claim 2 or a pharmaceutically acceptable salt thereof.

26. (Presently amended) A method of enhancing cognition in a healthy subject comprising administering a safe cognition enhancing amount of compound according to claim 1 or claim 2, or a pharmaceutically acceptable salt thereof.

27 to 34. (Canceled)